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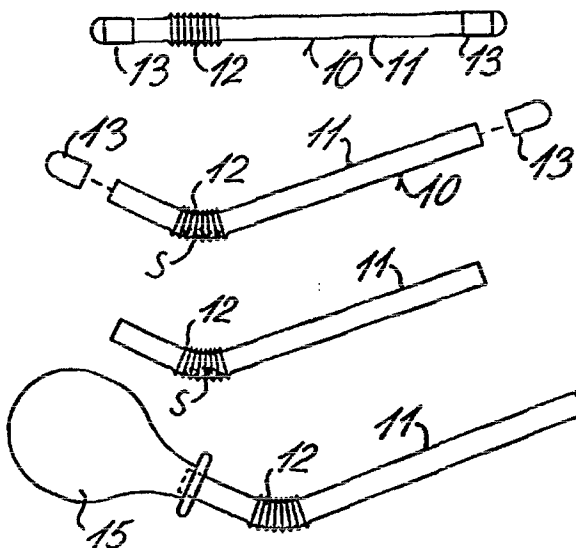
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(54) Title: AN INHALER

## (57) Abstract

An inhaler comprises a tubular body in which an air flow passage is defined. A single dose of an active, inhaleable, particulate substance is arranged within the air flow passage and is sealed or closed in relation to the ambient atmosphere by closure means, such as removable caps, a section of the flow passage extending from a free mouthpiece end of the tubular body along a major part of the total length of the flow passage is preferably 7-35 mm<sup>2</sup>, for example about 20 mm<sup>2</sup>. The inhaler may be adapted to be used only once, and the tubular body of the inhaler may be a length of a simple tube similar to a drinking straw.



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## AN INHALER

The present invention relates to an inhaler of the type comprising a tubular body defining an air flow passage therein.

5 Numerous inhalers of this type are known. As examples, US patent Nos. 4,524,769 and 4,907,583 and the international application WO 90/07351 disclose inhalers each comprising dosing means for supplying a dose of a particulate active substance to the air flow passage when the inhaler is to be  
10 used. The outlet end of the discharge passage is defined by a mouthpiece or nozzle which during use of the inhaler is positioned between the lips of the user. These known inhalers are of a relatively complicated structure, and they are not simple in use because the dosing means has to be operated  
15 each time the inhaler is to be used. Furthermore, these known devices are relatively heavy and bulky to carry in a pocket or in a ladies handbag.

When a particulate or powdered substance is inhaled through a mouthpiece or nozzle positioned between the users lips a  
20 substantial part of the active substance suspended in the inhaled air flow will not reach the lungs of the user, but may be swallowed or may impinge the mucous membrane of the oral cavity of the user. This does not only mean that a substantial amount of the active substance is lost, but also  
25 that the active substance coming into contact with the mucous membrane of the oral cavity or arriving at the stomach may have undesired side effects.

As disclosed in German Offenlegungsschrift No. 2815039 this problem may be solved by using a mouthpiece which in use  
30 extends into the oral cavity and along the tongue of the user. When a fluid flow including an active substance is directed into the oral cavity of the patient, for example from an inhaler or an atomizing device connected to the outer end of the mouthpiece, it is possible to direct such fluid

flow towards a desired part of the oral cavity, such as the throat or trachea of the patient.

The present invention provides an inhaler which may be produced in a very simple manner and at low cost and which is  
5 nevertheless very efficient.

The inhaler according to the invention comprises a tubular body defining an air flow passage therein and is characterized in comprising only a single dose of an active, inhalable, particulate substance arranged within the air flow  
10 passage, said dose being sealed or closed in relation to the ambient atmosphere by closure means which are to be removed or opened by a user prior to use, the inhaler being intended to be used only once. Thus, when the inhaler is to be used the closure means have to be removed or opened whereafter the  
15 active substance, which is positioned within the air flow passage, may be inhaled by the patient or user in a usual manner.

In the inhaler according to the invention the airflow passage may be without any tortuous sections which may promote precipitation or formation of a coating of the active substance  
20 on inner wall surface parts of the flow passage. The build up of such coating is further made unlikely when the inhaler is of a type being used only once. An inhaler which is used only once is much more hygienic than inhalers intended for multiple uses.  
25

The said closure means may be of any suitable type, such as caps, films, or foils which are removably arranged at the opposite ends of the tubular body. Alternatively, the closure means may comprise removable, puncturable or rupturable  
30 membranes extending across the air flow passage and being axially spaced therein so that the dose of active substance is arranged therebetween. The closure means could, of course, be of any other type, which is able to seal the dose of active substance in relation to the ambient atmosphere until

the inhaler is to be used, and which may be removed or opened by a patient or user prior to use.

According to another aspect the present invention provides an inhaler having a tubular body defining an air flow passage  
5 therein and means for supplying a dose of an active, inhale-able, particulate substance into the flow passage, and the inhaler according to the invention is characterized in that the cross-sectional area of the flow passage does not exceed  
10 75 mm<sup>2</sup> along the length of a flow passage section extending from a free mouthpiece end of the tubular body along a major part of the total length of the flow passage.

When such an inhaler is to be used by a patient a dose of an active substance is supplied into the flow passage by operating the supply means. The patient may now inhale the active  
15 substance by inserting the mouthpiece end between the lips and by forcefully inhaling air through the air flow passage defined in the inhaler. Because the cross-sectional area of the inner section of the air flow passage is relatively small, the air velocity in this section will be relatively  
20 high. Such high air velocity promotes atomization of the dose of particulate active substance and suspension of the finely divided particles in the air flow.

Also, when the inhaler has a single dose of active substance arranged within the air flow passage and is intended for  
25 single use only as described above, the cross-sectional area of the flow passage preferably does not exceed 75 mm<sup>2</sup> along the length of the flow passage section extending from a free mouthpiece of the tubular body along a major part of the total length of the flow passage.

30 In order to obtain a high air flow rate the cross-sectional area of the said flow passage section preferably does not exceed 70 mm<sup>2</sup> and is more preferably less than 50 mm. In the presently preferred embodiments the cross-sectional area of

the flow passage section is 7-35 mm<sup>2</sup> and preferably about 20 mm<sup>2</sup>.

The means for supplying a dose of an active substance into the flow passage may be of any known type to which doses of  
5 an active particulate substance may be fed in the form of small ampoules containing a single dose, or replaceable cartridges containing several doses. Alternatively, the inhaler may be adapted to be used only a few times or only once, and in such case the substance supply means may form a unitary  
10 part of the inhaler. As an example, the substance supply means may comprise one or a few dose containing chambers or pockets defined in the walls of the tubular body defining the air flow passage. Each dose containing chamber may be separated from the adjacent part of the flow passage by a film,  
15 membrane, or a wall part which may be punctured or ruptured by the user, for example by applying an inwardly directed pressure to the outer surface of the tubular body at the respective position.

The flow passage or the flow passage section may have any  
20 cross-sectional shape, such as square, rectangular, polygonal, elliptical, or circular. Furthermore, the cross-sectional area of the flow passage or the flow passage section may vary along the length thereof. Thus, the cross-sectional area of the flow passage or flow passage section  
25 may be smaller at the mouth piece end than at the opposite end of the flow passage. In the preferred embodiment, however, the flow passage or flow passage section has a substantially circular cross-section, the inner diameter of the flow passage section, which may be the total length of the air  
30 flow passage, being substantially the same along the length of the section. Thus, in a very simple embodiment the inhaler is formed similar to a drinking straw.

The tubular body of the inhaler and the air flow passage defined therein may have any suitable shape. Thus, the flow  
35 passage may have any suitable rectilinear, zigzag, angular or

curved course or any combination of such courses. Preferably, the flow passage comprises a curved section. As an example, the flow passage may comprise one or more pairs of substantially rectilinear sections and intermediate curved sections.

- 5 The tubular body of the inhaler may have a predetermined, permanent shape which cannot be changed by the user. In a preferred embodiment the tubular body comprises at least one bendable section so that the shape of the tubular body may be adapted to the form of the oral cavity of the individual
- 10 user. The walls of the tubular body forming the bendable section may be made from a deformable plastic material. Alternatively, the flexibility of the bendable section may be obtained by providing the bendable section of the tubular body with peripherally extending corrugations. The bottoms of
- 15 these corrugations may be provided with codes, such as colours, numbers, letters, or other kinds of indications, for assisting in obtaining a bend suitable for the individual user. When a patient or user has determined a shape of the tubular body which has been adapted to his individual oral
- 20 cavity, the user may read and note the visible code combination. When the user knows his individual code combination he may quickly adjust the bendable section of the tubular inhaler body next time he is using an inhaler of the same type.
- 25 In order to improve the dispersion of the particulate active substance in the air flowing through the air flow passage the inhaler may further comprise means for imparting a rotational movement to the air about the longitudinal axis of the flow passage. Such means may, for example, be helical grooves or
- 30 ribs formed on the wall surface parts of the tubular body defining the flow passage, or rotation imparting members arranged centrally within the air flow passage.

The tubular body of the inhaler may comprise a mouth piece section of any suitable length extending during use from the

35 teeth of the user to any desired position within the oral

cavity of the user. Thus, by suitably adapting the length and the shape of the mouthpiece section the inner end of the air flow passage may be positioned adjacent to and may be directed towards any part of the oral cavity which is to be treated by means of the active particulate substance. In cases where the active substance is to be inhaled to the lungs or bronchial tubes of the user or patient, for example when the active substance is a medicament for alleviating asthmatic deceases, the mouthpiece section of the tubular body preferably has a length so as to extend during use from the teeth of the user to a position adjacent to the root of the user's tongue. The mouthpiece section may then advantageously be shaped so that the inner end of the air flow passage is directed towards the throat or trachea inlet of the patient or user. Thereby it may be secured that substantially all of the active substance of the dose positioned in or supplied to the air flow passage will reach its target area.

In order to secure that the inhaler is correctly positioned within the oral cavity of the user or patient the inhaler may further comprise a bite piece formed on the outer surface of the tubular body for engaging with the upper jaw teeth of the user. Such bite piece may form an integral part of the tubular body. It is preferred, however, that the shape of the bite piece is adapted to the teeth of the individual user. In such case the relatively expensive and individually formed bite piece is removably mounted on the tubular body of the inhaler so that the bite piece may be kept and reused when the tubular body is discarded. Such removably mounted bite piece may advantageously be used in connection with tubular bodies adapted to be used only a few times or only once. The bite piece may be combined with a set of false teeth. Thus, such a set of false teeth may comprise means for positioning the tubular body in relation to the oral cavity of the user, or the bite piece may be individually shaped when the false teeth are made to the individual user.



The tubular body of the inhaler may have a fixed length. Alternatively, the tubular body may be movable from a retracted storage condition to an extended condition of use. This may, for example, be obtained by providing the tubular  
5 body with peripheral corrugations along a major part of its length so as to allow longitudinal stretching of the tubular body. As another possibility, the tubular body may comprise telescopically cooperating tubular parts allowing such parts to be moved between retracted and extended positions. The  
10 closure means sealing the dose of active substance in relation to the ambient atmosphere or the supply means for supplying such dose into the air flow passage of the inhaler may be opened or actuated automatically when the tubular body is moved from its retracted storage condition to its extended  
15 condition of use.

In order to completely eliminate the risk that larger particles of the active substance or of foreign matter, such as ruptured or removed closing means, are inhaled by a patient the inhaler may further comprise retaining means arranged  
20 within the flow passage for retaining particles of a size exceeding a predetermined size. Such retaining means may, for example, comprise a sieve or screen extending across the flow passage and being arranged downstream of the dose of active substance to be inhaled. Alternatively, when the closure  
25 means comprise removable closure caps, such caps may be integrally connected to the tubular body by flexible strings or bands. Alternatively or additionally, the free end portions of the tubular body may be bent towards each other so as to position the free ends closely adjacent, and the free  
30 ends may then be closed by a common, removable closure member, such as a pair of interconnected or integrally formed closure caps. Furthermore, at least one of the cap members is preferably made from a transparent material, or the tubular body is at least partly made from a transparent material, so  
35 that the patient or user may see that the tubular body contains a dose of the active substance.

In most cases the patient or user is able to inhale atmospheric air through the air flow passage so as to obtain an air flow being sufficiently vigorous to suspend the active substance therein in an atomized condition. However, smaller  
5 children and adult persons having a substantially reduced lung capacity may not be able to generate a sufficient air flow rate through the air flow passage. Therefore, the inhaler may further comprise forced flow generating means for providing a forced air flow through the flow passage. Such  
10 flow generating means may be of any kind which is able to generate pressurized air. As an example, the forced flow generating means may comprise a compressible bulb to be mounted on the outer end or air inlet end of the tubular body. The flow generating means may then be actuated at the  
15 same time as the patient is inhaling.

It should be understood that the inhaler according to the present invention should not necessarily be inserted into the oral cavity of the user, and the term "mouthpiece" is not intended to indicate that it should necessarily be inserted  
20 into the user's mouth. Thus, one end or the mouthpiece end of the tubular body may be adapted to be inserted into a nostril of a user or patient. In such case, the inhaler preferably comprises a pair of tubular bodies and a connecting part for interconnecting the same, said one end or mouthpiece end of  
25 said pair of tubular bodies being arranged in spaced relationship, so that said ends may be inserted into the nostrils of a user or patient.

The invention further provides substance supply means for use in connection with an inhaler of the type described above  
30 including such supply means, said supply means comprising a tube length containing only a single dose of a particulate active substance, said dose being sealed or closed in relation to the ambient atmosphere by closure means which are to be removed or opened by a user prior to use, one end of the  
35 tubular length being insertable in or connectable to the air flow passage of the inhaler. The tube length which forms a capsule for the active substance is preferably bendable, for

example due to a plurality of adjacent peripherally extending corrugations. The tube length may be closed at opposite ends by means of removable closure caps. The free ends of the tube length may be bent together and be closed by a common closure  
5 member or interconnected closure caps.

The invention will now be further described with reference to the drawings, wherein

- Fig. 1 illustrates a first embodiment of the inhaler according to the invention,
- 10 Fig. 2 illustrates a second embodiment of the inhaler according to the invention,
- Fig. 3 illustrates the use of the inhaler shown in Fig. 1,
- Fig. 4 illustrates a third embodiment of the inhaler according to the invention,
- 15 Fig. 5 illustrates a fourth embodiment of the inhaler according to the invention,
- Fig. 6 illustrates how the inhaler shown in Fig. 5 may be used,
- Fig. 7 illustrates how a bent portion of the inhaler shown in  
20 Fig. 6 may be provided with bending codes,
- Fig. 8 illustrates a fifth embodiment of the inhaler according to the invention,
- Fig. 9 illustrates how the inhaler shown in Fig. 8 may be used,
- 25 Fig. 10 illustrates a mouthpiece for a sixth embodiment of the inhaler according to the invention,
- Fig. 11 illustrates how the mouthpiece shown in Fig. 10 may be positioned in the oral cavity of the user,
- Figs. 12 and 13 illustrate in an enlarged scale two different  
30 devices for supplying separate doses of active substance to the flow passage of the mouthpiece shown in Figs. 10 and 11,
- Fig. 14 illustrates the mouthpiece shown in Fig. 10 provided with a third device for supplying doses of active substance,
- Fig. 15 illustrates a further embodiment of a mouthpiece for  
35 use in connection with the inhaler according to the invention,

Fig. 16 illustrates the mouthpieces shown in Fig. 15 provided with a device for supplying active substance thereto, Fig. 17 illustrates the function of the inhaler shown in Figs. 15 and 16,

5 Fig. 18 illustrates a seventh embodiment of the inhaler according to the invention,

Fig. 19 illustrates an eighth embodiment of the inhaler according to the invention,

10 Fig. 20. illustrates how the inhaler shown in Fig. 19 may be used,

Fig. 21 shows a capsule containing a single dose of an active substance,

Fig. 22 illustrates how the capsule shown in Fig. 21 may be used in connection with a mouthpiece as shown in Figs. 14-16

15 Fig. 23 shows a ninth embodiment of the inhaler, and

Fig. 24 shows a tenth embodiment, in which the opposite ends of the tubular inhaler is closed by a common closure cap.

Fig. 1 shows an inhaler 10. When delivered from the manufacturer, the inhaler may comprise a straight, thin-walled  
20 tubular body 11 having a bendable section 12 and removable caps 13 closing the opposite open ends of the tubular body 11. The inner bore of the tubular body 11 which defines an air flow passage of the inhaler contains a single dose of a particulate or powdered active substance, such as steroids,  
25  $\beta_2$ -agonists, anticholinergica, or other medical products. The tubular body 11 may have a circular cross-sectional shape and have a substantially uniform inner diameter and wall thickness along the length of the tubular body and may be similar to a drinking straw. The section 12 may have peripheral  
30 corrugations so as to be bendable. The tubular body 11 may, for example, be made from a suitable plastic material by extrusion, and the inner diameter of the tubular body is preferably within the range of 4-8 mm, for example 5-6 mm. The material of the tubular body 11 may have been treated so  
35 as to reduce or eliminate the possibility of static electricity.

The inhaler 10 shown in Fig. 1a which contains only a single dose of the active substances is intended to be used only once whereafter the inhaler is discarded. A suitable small number of disposable inhalers of this type may be packed, for example similar to cigarettes, and they may be carried by the user in a pocket or a ladies handbag without occupying much space.

When an inhaler 10 of the type shown in Fig. 1a is to be used the user or patient may shake the inhaler so as to disintegrate the particulate active substances contained therein. The bendable section 12 may now be bent whereby the active substance S is positioned within the corrugation troughs of the bendable section 12 as indicated in Figs. 1a and 1b. Thereafter the caps 13 may be removed as shown in Fig. 1b. The inhaler 10 is then ready for use, and a longer straight end portion of the tubular body may be inserted into the oral cavity 14 of a user or patient as shown in Fig. 3. Because the tubular body 11 has been bent the active substance S contained in the corrugation troughs of the section 12 may be prevented from falling out from the tubular body.

As shown in Fig. 3 the inner end of the inhaler is positioned adjacent to the root of the patient's tongue. When the patient inhales air through the air flow passage defined within the tubular body 11 the particulate active substance S is withdrawn from the corrugation troughs of the section 12 and is suspended in the air flow which is inhaled into the patient's lungs. In case the patient is a small child or for some other reason is not able to inhale sufficiently vigorously a compressible bulb 15 or other means for generating a forced flow through the tubular body 11 may be mounted on the outer end thereof as shown in Fig. 1d. A flow of air with active substance suspended therein may then be blown into the oral cavity 14 at the same time as when the patient inhales. The embodiment shown in Fig. 2 is slightly modified in relation to the embodiment shown in Fig. 1. Thus, in Fig. 2 the tubular body 11 is provided with a helically extending corru-

gation 16 which may impart a rotational movement to the air flow being inhaled through the air flow passage defined in the tubular body 11. The tubular body 11 and/or at least one of the closure caps 13 is preferably made from a transparent material so that the user may visually make sure that the inhaler contains a dose of an active substance.

Fig. 4 shows an embodiment 10 having not only a bendable section 12 provided with peripherally extending corrugations, but also a bendable section 17 without such corrugations at an opposite end portion of the tubular body 11. In fact, the material and the wall thickness of the tubular body 11 may be chosen so that any part of the tubular body may be bent into a desired shape. The inhaler shown in Fig. 5 corresponds to that shown in Fig. 4. The only difference is that the embodiment shown in Fig. 5 comprises a bendable section 12 at opposite end portions of the tubular body, which sections 12 both comprise peripherally extending corrugations.

For some patients who are not sensing so well it may be difficult to immediately position the inhalers shown in Figs. 1, 2, 4, and 5 correctly in the oral cavity 14. Therefore, the tubular body 11 may advantageously be inserted in a bite piece or teeth block 18 of the type shown in Fig. 5. Fig 5d is an end view of the bite piece 18, while Fig. 5e is a longitudinally sectional view of the bite piece. The bite piece 18 comprises a longitudinally extending channel or slot 19 which is dimensioned so that the tubular body 11 may be snugly received therein as indicated by an arrow in Fig. 5d. Troughs or grooves 20 and 21 are defined in the upper outer surface of the bite piece 18, and a trough 22 is defined in the lower outer surface of the bite piece. When the bite piece or teeth block 18 has been mounted on the tubular body 11 which has been bent into the desired shape and the closure caps 13 have been removed as explained above, the inhaler assembly comprising the tubular body 11 and the bite piece 18 may be inserted into the user's mouth. As shown in Fig. 6 the bite piece or teeth block 18 may then be positioned so that

the upper lip 23 and the upper teeth 24 of the patient are positioned in the troughs 20 and 21, respectively, while the lower lip 25 of the user is positioned in the trough 22 of the bite piece or teeth block 18 whereby the tubular body 11  
5 may be positioned very accurately within the user's oral cavity 14.

As shown in Fig. 6 the inner end portion of the tubular body 11 may be shaped so that the inner open end of the tubular body is positioned adjacent to and directed towards the  
10 throat 26 of the user or patient. Thus, almost all of the active substance contained within the tubular body may be transferred to the patient's lungs when the patient vigorously inhales air through the air flow passage defined within the tubular body 11. It should be understood that the inner  
15 end portion of the tubular body 11 could be directed towards any desired surface part of the oral cavity to be treated by the active substance contained in the inhaler.

In order to ensure that the open inner end of the tubular body 11 is directed towards the throat 26 of the patient or user as shown in Fig. 6, or towards any other surface part of the oral cavity 14 to be treated it is important that the  
20 shape of the inner end of the tubular body 11 is adapted to each individual user. When the inner end of the tubular body has a bendable section 12 provided with peripherally extending corrugations, the bend or curvature which is adapted to  
25 the oral cavity of a specific patient or user may be expressed as a code which may be remembered. Fig. 7 illustrates examples of such coding. As indicated in Fig. 7a a number may be assigned to each or every second of the peripheral troughs formed between the adjacent peripheral corrugations of the bendable section 12. The number code to be  
30 remembered by the user may then indicate which of the upwardly facing trough parts should be fully opened and which should not.

Alternatively, adjacent troughs may be differently coloured or different colours may otherwise be assigned to the various troughs as indicated in Figs. 7b and 7c. The bend or curvature suitable for each individual user or patient may then be  
5 expressed as a colour code in a similar manner as explained above in connection with the number coding.

The inhaler 10 illustrated in Fig. 8 is of the same type as that described above with reference to Fig. 2. However, the inner end portion of the tubular body 11 shown in Fig. 8 has  
10 a permanent bend 27 which is made when the inhaler is being manufactured.

Like in the inhaler shown in Fig. 2 the tubular body 11 has a helical corrugation 16 extending along the length of the tubular body. As explained above in connection with Fig. 2,  
15 such helical corrugation may tend to impart a rotational movement to air inhaled through the tubular body 11. The tubular body 11 in any of the embodiments shown in Figs. 1, 2, 4, 5 and 8 may, for example, be made by injection moulding, blow moulding, or extrusion. In the latter case, the  
20 corrugations may be formed in the walls of the tubular body 11 during a subsequent manufacturing stage. The tubular bodies 11 shown in Figs. 2 and 8 may, alternatively, be made by helically winding a strip of sheet material, such as paper, paperboard or another fibrous sheet material, with  
25 mutually overlapping adjacent edge portions. Such edge portions may then be interconnected or sealed, whereby the helically extending corrugation 16 may be formed. In order to facilitate mounting of the snugly fitting caps 13 on the opposite ends of the tubular body 11 when a dose of active  
30 material has been arranged therein, the tubular bodies 11 illustrated in Figs. 1, 2, 4, 5, and 8 may be cut on the bias as indicated with dotted lines at 28 in Fig. 8c.

The tubular body 11 shown in Fig. 8 may be inserted into the channel or slot 19 of a bite piece or teeth block 18 as  
35 explained above in connection with Fig. 5. Thereafter, when



the bendable section 12 has been given the desired shape and the caps 13 have been removed, the inhaler may be positioned in the mouth of a patient as illustrated in Fig. 9 and as explained above in connection with Fig. 6.

5 Fig. 10 illustrates an embodiment of a tubular body 29 for an inhaler according to the invention. Figs. 10a, 10b, 10c, and 10d illustrate a longitudinally sectional view, a top plan view, an end view, and a cross-sectional view along the line D-D, respectively. The tubular body 29 defines a longitudi-  
10 nally extending air flow passage 30 having a substantially uniform cross-sectional area along the length thereof. The outer end (the left hand end in Fig. 10) of the tubular body 29 has a shape corresponding to the shape of the bite piece or teeth block 18 shown in Figs. 5 and 8. This means that in  
15 the embodiment shown in Fig. 10 the bite block is formed integrally with the tubular body 29. Thus, the tubular body 29 has troughs or grooves 20 and 21 formed in the upper outer surface of the outer end portion of the tubular body 29. These troughs 20 and 21 are intended to receive the upper lip  
20 23 and the upper teeth 24, respectively, of a user or patient. Furthermore, a trough 22 is formed in the lower outer surface of the tubular body 29 for receiving the lower lip 25 of the user as illustrated in Fig. 11.

The tubular body 29 illustrated in Fig. 10, which is preferably made from plastic by injection moulding, may be adapted  
25 to be used only once. In such case, a single dose of an active powdered or particulate material may be arranged within the air flow passage 30, and the open ends of the air flow passage may be sealed or closed by removable sealing or  
30 closing means, such as a film or foil which may be torn off. Alternatively, the tubular body 29 illustrated in Fig. 10 may be adapted to cooperate with a dose feeding device for feeding a dose of powdered or particulate active material into the air flow passage 30 when the inhaler is to be used. In  
35 that case the tubular body 29 is preferably adapted to be

used several times, and each sample may then have a shape which has been adapted to the individual user.

Figs. 12 and 13 show the outer end portion or bite piece of the tubular body 29 provided with a separate dose feeding device 31. The dose feeding device 31 comprises a short tube section 32 which may be inserted into the air flow passage 30 of the tubular body 29. In Fig. 12 the dose feeding device 31 comprises a disc 33 which is rotatably mounted within a housing 34. The disc 33 comprises a plurality of capsules 35 in a circular arrangement. The capsules 35 of the disc 33 may successively be indexed to a position of use. By operating parts of the housing 34 the opposite ends 36 of the capsule 35 in this position are cut off whereby the remaining tubular part of the capsule is brought into communication with the tube section 32 and the air flow passage 30 defined in the tubular body 29. The dose of active substance contained in the capsule being cut may now be inhaled when the tubular body 29 has been inserted into the oral cavity of a patient as illustrated in Fig. 11.

The dose feeding device 31 illustrated in Fig. 13 is similar to that illustrated in Fig. 12. In Fig. 13, however, the capsule carrying disc 33 has been replaced by a strip 37 carrying a plurality of capsules 35 arranged in a rectilinear row. In other respects, the dose feeding device 31 of Fig. 13 functions substantially in the same manner as the feeding device shown in Fig. 12. In order to prevent inhalation of the cut capsule ends 36 or other foreign matter, a sieve or grid 38 allowing the particulate active substance, but not the capsule ends 36, to pass may be arranged within the air flow passage 30 as indicated in Figs. 12 and 13.

Fig. 14 illustrates an embodiment of the inhaler according to the invention comprising a tubular body 29 as that illustrated in Fig. 10. The inhaler further comprises a small container or capsule 39 which may contain a single dose of a particulate or pulverulent active substance. Fig. 14a is an

end view of the inhaler, while Fig. 14b illustrates the inhaler in a side view and partially sectional view. The capsule 39 comprises a tube stub or tube section 32 which may be inserted into the outer end of the air flow passage 30, which means that the capsule 39 is replaceable. When the inhaler is to be used closure means (not shown) of the capsule 39 may be removed or ruptured to allow air to flow through the capsule and into the air flow passage 30. As an example, such closure means may comprise a removable cap or a film for closing the tube stub 32 and a removable film or foil closing an opening in the capsule 39, or a removable wall part of the capsule. The capsule or container 39 may then be disposed of when it has been used a single time while the tubular body 29 may be used several times.

Fig. 15 illustrates an alternative embodiment of the tubular body shown in Fig. 14. Figs. 15a, 15b, and 15c are an end view, a longitudinally sectional view, and a cross-sectional view along the line C-C, respectively. It should be noted that Fig. 15c has been shown in a larger scale than Figs. 15a and 15b. In addition to the air flow passage 30 the tubular body 29 shown in Fig. 15 comprises longitudinally extending, through-going upper and lower air passages 40 and 41, respectively. When the tubular body 29 is being used and a flow of air with active substance dispersed therein is being inhaled through the air flow passage 30, flows of "false air" are simultaneously being inhaled through the air passages 40 and 41. These flows of false air envelope the air flow in which the active substance is dispersed so as to direct the dispersed active substance in the desired direction and so as to reduce loss of active substance.

Fig. 16 illustrates an inhaler embodiment comprising a tubular body 29 as that shown in Fig. 15 and a replaceable container or capsule 39 of the type which is shown in Fig. 14 and which is replaceably mounted on the tubular body 29 by means of a tube stub 32. The capsule 39 is shaped so that it does not close the inlet openings of the air passages 40 and

41. Fig. 16a is an end view of the inhaler while Fig. 16b is a side view and partially sectional view of the inhaler. Apart from the enveloping air flowing through the air passages 40 and 41 the inhaler illustrated in Fig. 16 may be  
5 operated and may function substantially as described above with reference to Figs. 14 and 15.

Fig. 17 diagrammatically illustrates the inner end of the inhaler shown in Fig. 16 which has been inserted into the oral cavity 14 of a user or patient. When the patient inhales  
10 air through the inhaler an air flow 42 with active substance dispersed therein will be enveloped by flows of atmospheric air which are represented by arrows 43 and 44 and which contain no or few particles of active substance. Such enveloping air flows may assist in conveying the dispersed active  
15 substance into the patient's lungs without any substantial loss of active substance.

Fig. 18 illustrates a further embodiment of the tubular body 29. Fig. 18a is an end view and Fig. 18b is a longitudinally sectional view of the tubular body 29 while Fig. 18c is an  
20 end view in the direction C-C shown in an enlarged scale. The tubular body 29 shown in Fig. 18 is a short version having a shape similar to the outer end portion of the tubular body shown in Fig. 15b at the left hand side of the section line C-C. However, as best illustrated in Fig. 18a the outer end  
25 portion of the air flow passage 30 may be in the form of a longitudinally extending channel or groove 45. The tubular body 29 shown in Fig. 18 may be used in connection with a capsule or container 39 as shown in Figs. 14 and 16, and the tube stub 32 of the capsule 39 may then be received in the  
30 channel or groove 45, and the axial length of the channel or groove may correspond to the length of the tube stub or tube section 32 of the capsule 39.

In Fig. 19 an embodiment corresponding to that of Fig. 1 has been shown. However, in addition to the bendable section 12  
35 the tubular body 11 has a second similar bendable section 46

which is spaced from the bendable section 12 by a non-corrugated, rectilinear tubular section 47. The tubular body 11 contains a single dose of an active substance S and when the inhaler 10 is to be used the sections 12 and 46 may be bent as shown in Figs. 19b and 19c so that the sections 12, 47 and 46 are substantially S-shaped. The active substance S is mainly received in the inner corrugation troughs of the bendable tubular section 12. Now, the removable closure caps 13 may be removed from the opposite ends of the tubular body 11 as illustrated in Fig. 19c, and the tubular body may be inserted into the oral cavity 14 of a user as illustrated in Fig. 20. Even when the user reclines his head as shown in Fig. 20 the active substance S may remain in the inner corrugation troughs of the bendable section 12. When, however, the user or patient inhales air through the air flow passage of the tubular body 11, the velocity of flowing air causes a static pressure drop so that the active substance S is sucked from the corrugation troughs and entrained with and efficiently dispersed in the air flow.

If an inhaler as that shown in Figs. 1, 2, and 4 is shortened it may be used as a disposable container or capsule for a single dose of an active substance S. Such a tubular container or capsule is shown in Fig. 21 and may be closed at its opposite ends by removable closure caps 13 or by any other removable or breakable closure means. The tubular capsule may have a central bendable section 49 having peripheral corrugations as those previously described. The tubular container or capsule 48 may be used together with any of the tubular bodies 29 shown in Figs. 10-18. Fig. 22 illustrates how the container or capsule 48 may be used in connection with the tubular body 29 shown in Figs. 15 and 16. When the tubular capsule 48 is to be used it may be bent as illustrated in Fig. 21b so that the active substance S is collected in the bendable section 49 and is mainly received in the inner corrugation troughs defined therein. When the closure caps 13 have been removed, one end of the capsule 48 may be inserted into the outer end of the air flow passage

of the tubular body 29 as illustrated in Fig. 22. Now, the inhaler is ready for use in a manner previously described. When the active substance S from the capsule 48 has been inhaled, the capsule may be discarded and a new capsule is  
5 used for the next inhalation.

Fig. 23 shows a tubular inhaler 10 in which the section 12 has a number of annular corrugations. Figs. 23a and 23b illustrate the rectilinear inhaler prior to use and the bent inhaler made ready for use, respectively. As shown in Fig.  
10 23c the corrugations are of a type which is substantially saw tooth shaped in an axial sectional view. Thus, the valleys 50 and peaks 51 of the corrugations are relatively sharp.

In the embodiment shown in Fig. 24 the tubular inhaler 10 is stored in a condition in which the inhaler 10 is bent to a  
15 position in which the free ends of the inhaler are positioned closely adjacent, and the inhaler with the active substance therein is maintained in this position by means of a single closure member 52 which is closing both of the free ends of the inhaler. The closure member may, for example, be in the  
20 form of a pair of closure caps which are interconnected by a connecting portion which may be formed integrally with the closure caps. This embodiment secures that both of the opposite ends of the tubular inhaler 10 are opened before the inhaler can be used.

25 It should be understood that various amendments and modifications of the embodiments described above and shown in the drawings could be made without departing from the scope of the present invention. Thus, features described in connection with any of the embodiments shown could also be used in  
30 connection with one or more of the other embodiments shown. As an example, any of the embodiments may contain a sieve or grid 38 as shown in Figs. 12 and 13 or other kinds of means for retaining larger particles. Furthermore, any of the tubular bodies shown may comprise separate passages for  
35 "false" or enveloping air. Similarly, any of the embodiments

of the tubular body may be provided with a compressible bulb 15 as shown in Fig. 1d or any other kind of means for providing compressed air.

## CLAIMS

1. An inhaler comprising a tubular body (11) defining an air flow passage therein,  
characterized in comprising only a single dose of an active,  
5 inhalable, particulate substance (S) arranged within the air flow passage, said dose being sealed or closed in relation to the ambient atmosphere by closure means (13) which are to be removed or opened by a user prior to use, the inhaler being intended to be used only once.
- 10 2. An inhaler having a tubular body (11, 29) defining an air flow passage (30) therein and means (31, 39, 48) for supplying a dose of an active, inhalable, particulate substance (S) into the flow passage (30),  
c h a r a c t e r i z e d in that the cross-sectional area  
15 of the flow passage does not exceed  $75 \text{ mm}^2$  along the length of a flow passage section extending from a free mouthpiece end of the tubular body (30) along a major part of the total length of the flow passage.
- 20 3. An inhaler according to claim 1, wherein the cross-sectional area of the flow passage does not exceed  $75 \text{ mm}^2$  along the length of a flow passage section extending from a free mouthpiece end of the tubular body along a major part of the total length of the flow passage.
- 25 4. An inhaler according to claim 2 or 3, wherein the cross-sectional area of the flow passage section (39) does not exceed  $70 \text{ mm}^2$  and is preferably less than  $50 \text{ mm}^2$ .
5. An inhaler according to claim 4, wherein the cross-sectional area of the flow passage section (39) is  $7\text{-}35 \text{ mm}^2$ , preferably about  $20 \text{ mm}^2$ .
- 30 6. An inhaler according to claim 4 or 5 comprising only a few or a single dose of the active substance (S) and being of a type adapted to be used only a few times or once.



7. An inhaler according to any of the claims 1-6, wherein the flow passage (39) or flow passage section has a substantially circular cross-section, the inner diameter of the flow passage being substantially the same along the length of the section.
8. An inhaler according to claim 7, wherein the inhaler is formed similar to a drinking straw.
9. An inhaler according to any of the claims 1-8, wherein the flow passage comprises a curved section (27).
10. An inhaler according to any of the claims 1-9, wherein the tubular body comprises a bendable section (12, 17).
11. An inhaler according to claim 10, wherein the bendable section (12, 17) of the tubular body (11) comprises peripherally extending corrugations.
12. An inhaler according to claim 11, wherein the bottoms of the corrugations are provided with codes for assisting in obtaining a bend suitable for the individual user.
13. An inhaler according to any of the claims 1-12, further comprising means (16) for imparting to air flowing through the flow passage a rotational movement about the longitudinal axis of the flow passage.
14. An inhaler according to any of the claims 1-13, wherein the tubular body (11, 29) comprises a mouthpiece section having a length so as to extend during use from the teeth (24) of the user to a position adjacent to the root of the user's tongue.
15. An inhaler according to any of the claims 1-14, further comprising a bite piece (18) formed on the outer surface of the tubular body (11, 29) for engaging with the upper jaw

teeth (24) of the user so as to position the inhaler in the oral cavity (14) of the user.

16. An inhaler according to claim 15, wherein the bite piece (18) is removably mounted on the tubular body (11).

5 17. An inhaler according to claim 15 or 16, wherein the shape of the bite piece (18) is adapted to the teeth (24) of the individual user.

18. An inhaler according to any of the claims 1-17, wherein the tubular body is moveable from a retracted storage condi-  
10 tion to an extended condition of use.

19. An inhaler according to claim 18, wherein the tubular body is provided with peripheral corrugations along a major part of its length so as to allow longitudinal stretching of the tubular body.

15 20. An inhaler according to claim 18, wherein the tubular body comprises telescopically cooperating tubular parts.

21. An inhaler according to any of the claims 1-20, further comprising retaining means (38) arranged within the flow passage (30) for retaining particles of a size exceeding a  
20 predetermined size.

22. An inhaler according to claim 21, wherein the retaining means comprises a sieve or screen (30) extending across the flow passage (30).

23. An inhaler according to any of the claims 1 and 3-22,  
25 wherein the closure means comprise a pair of cap members (13) removably mounted at opposite ends of the tubular body (11).

24. An inhaler according to any of the claims 10-23, wherein the free ends of the tubular body (11) are positioned closely

adjacent, said free ends being closed by a common removable closure member.

25. An inhaler according to claim 23 or 24, wherein at least one of the cap members or closure members (13) is made from a transparent material.

26. An inhaler according to any of the claims 1-25, wherein the tubular body (11, 29) is at least partly made from a transparent material.

27. An inhaler according to any of the claims 1-26, further comprising forced flow generating means (15) for providing a forced air flow through the flow passage.

28. An inhaler according to claim 27, wherein the forced flow generating means comprises a compressible bulb (15) to be mounted on an air inlet end of the tubular body (11).

29. An inhaler according to any of the claims 1-28, wherein one end or the mouthpiece end of the tubular body is adapted to be inserted into a nostril of a user.

30. An inhaler according to claim 29 comprising a pair of tubular bodies and a connecting part for interconnecting the same, said one end or mouthpiece end of said pair of tubular bodies being arranged in spaced relationship so that said ends may be inserted into the nostrils of a user.

31. A substance supply means for use in connection with an inhaler according to any of the claims 2-30, said supply means comprising a tube length (48) containing only a single dose of a particulate active substance (S), said dose being sealed or closed in relation to the ambient atmosphere by closure means (13) which are to be removed or opened by a user prior to use, one end of the tubular length being insertable in or connectable to the air flow passage (30) of the inhaler.

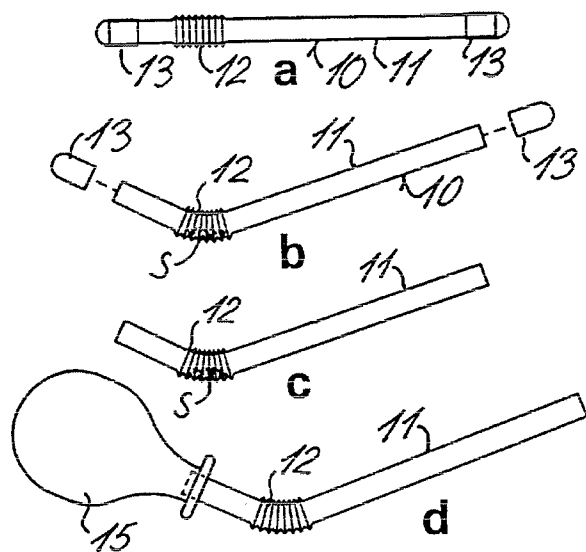
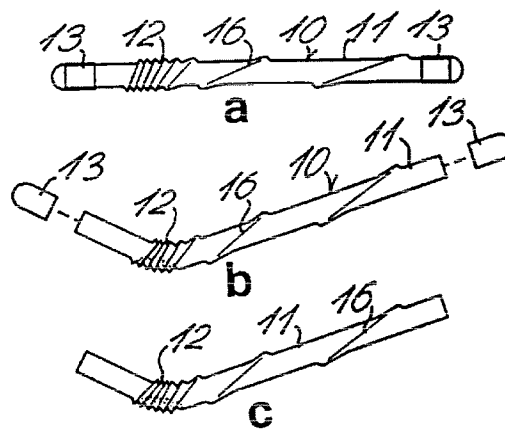
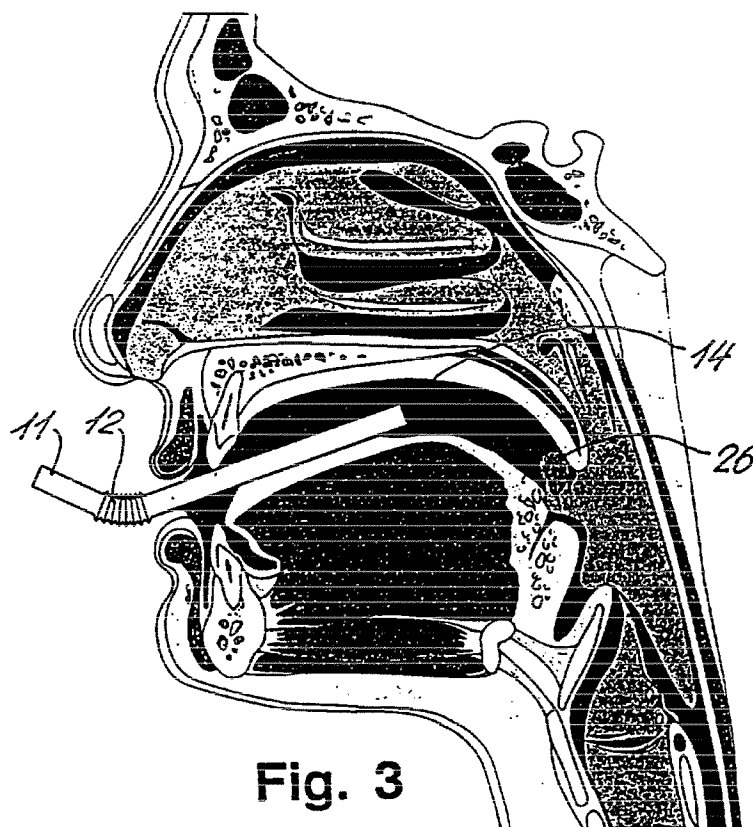
32. A substance supply means according to claim 31, wherein the tubular length (48) comprises at least one bendable section (49).

33. A substance supply means according to claim 32, wherein  
5 the bendable section comprises a plurality of adjacent, peripherally extending corrugations.

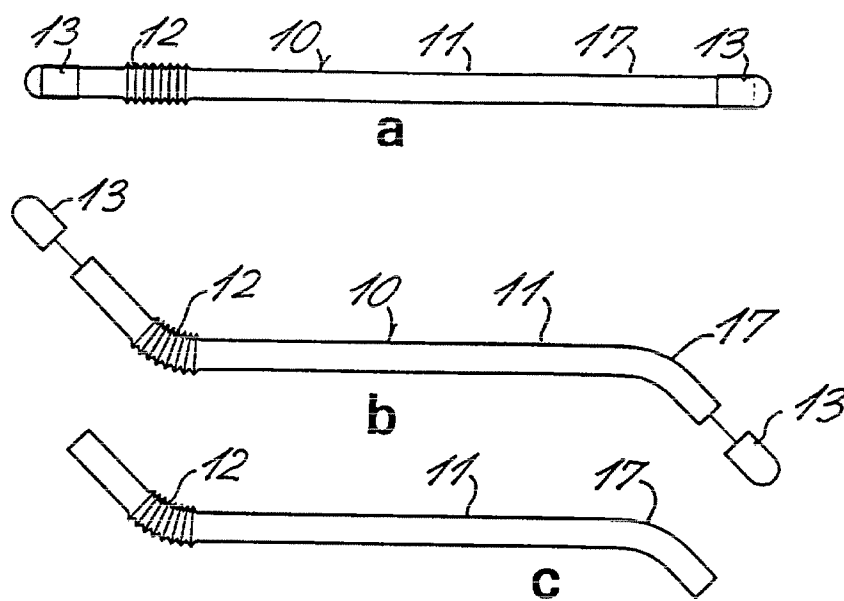
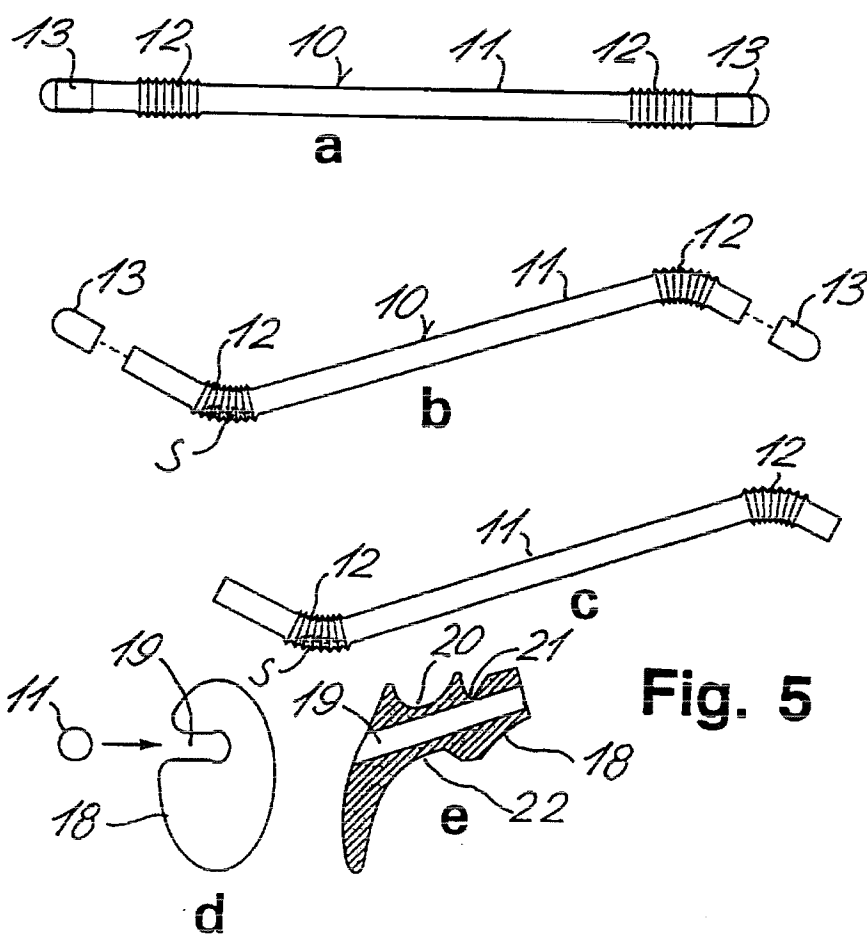
34. A substance supply means according to claim 32 or 33, wherein the closure means comprise a pair of removable closure caps (13) closing opposite ends of the tube length (48).

10 35. A substance supply means according to claim 32 or 33, wherein the free ends of the tube length (48) are positioned closely adjacent, said free ends being closed by a common removable closure member.

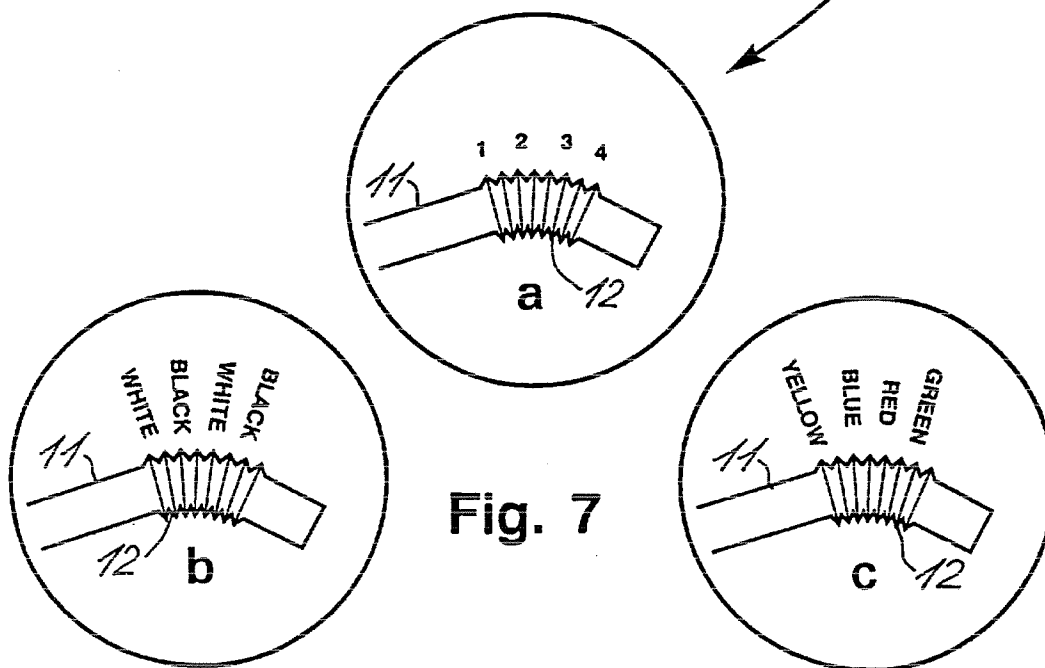
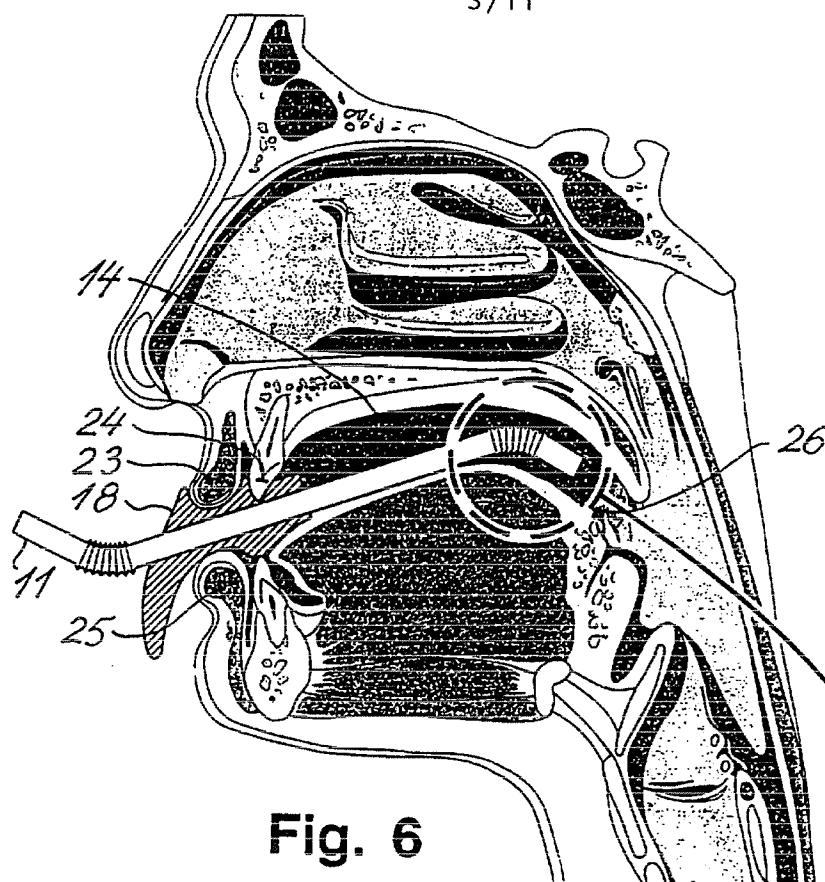
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**Fig. 1****Fig. 2****Fig. 3**

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**Fig. 4****Fig. 5**

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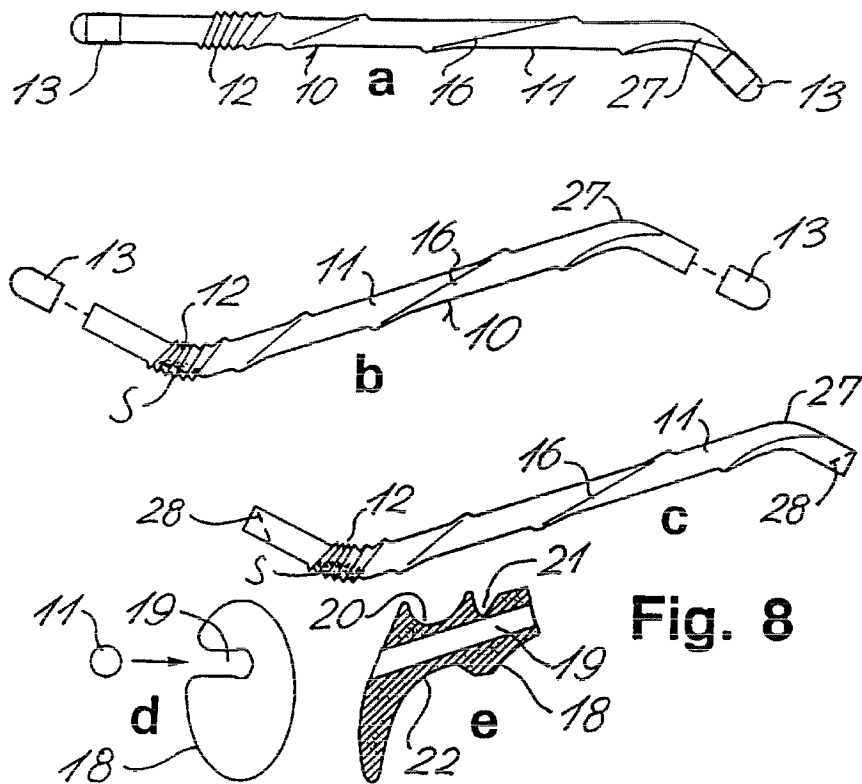


Fig. 8

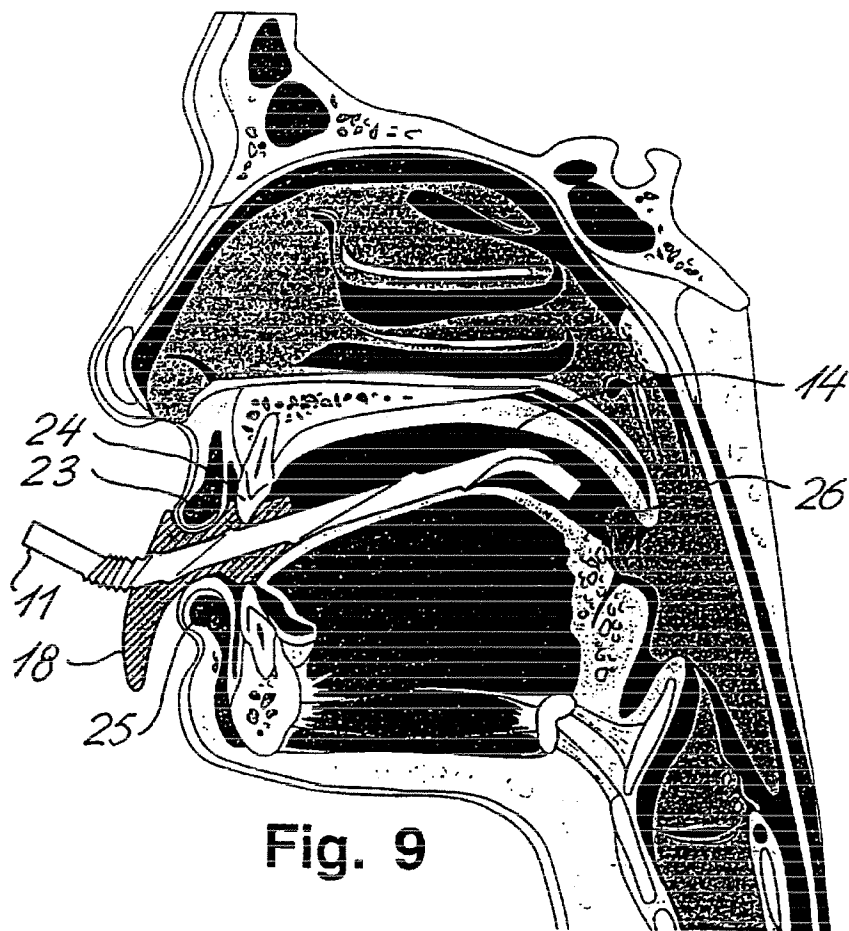
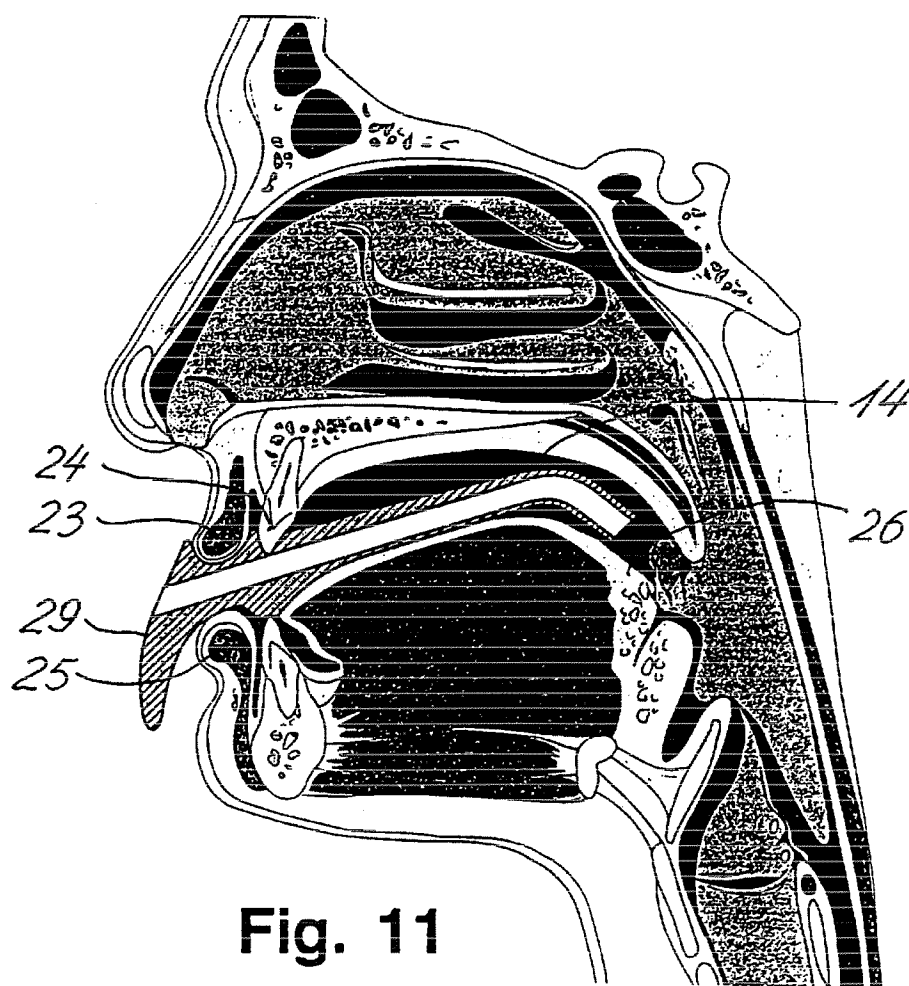
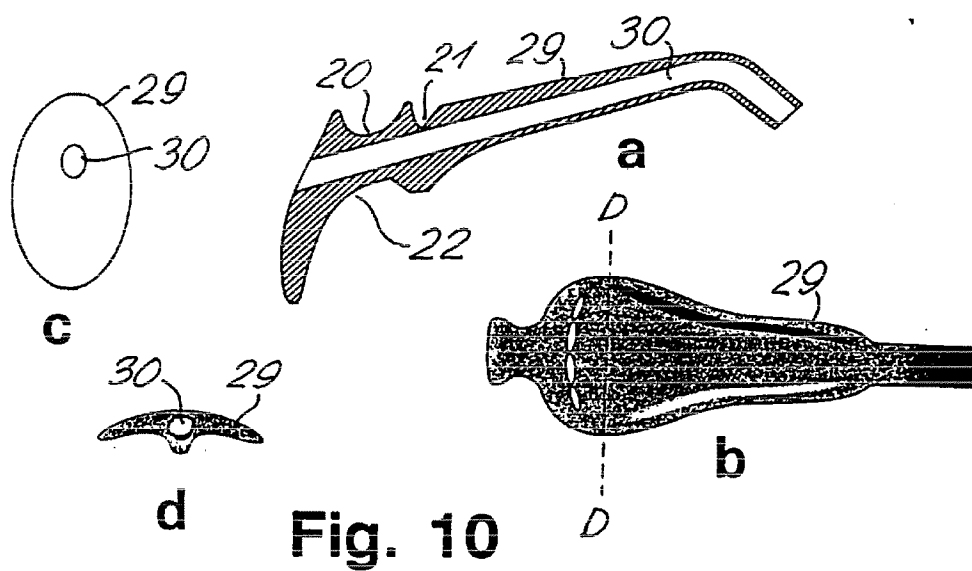


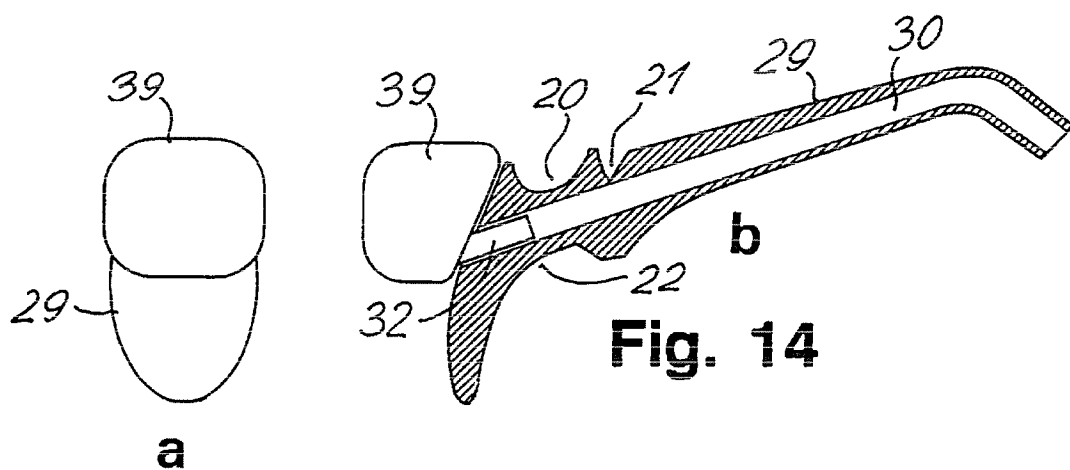
Fig. 9



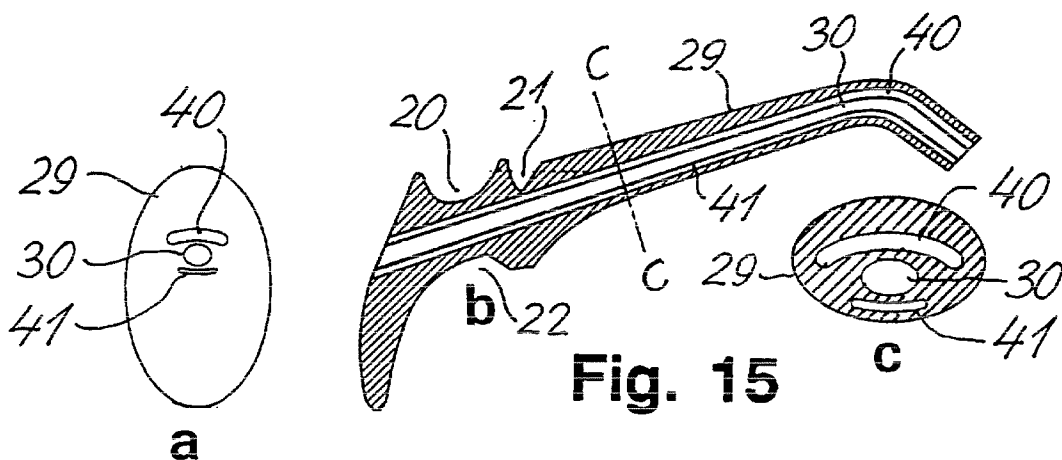




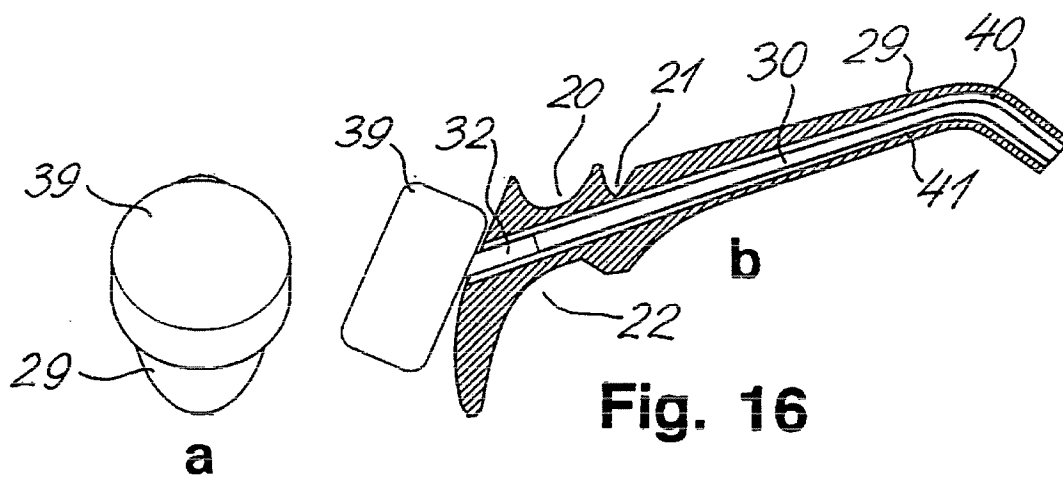
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**Fig. 14**

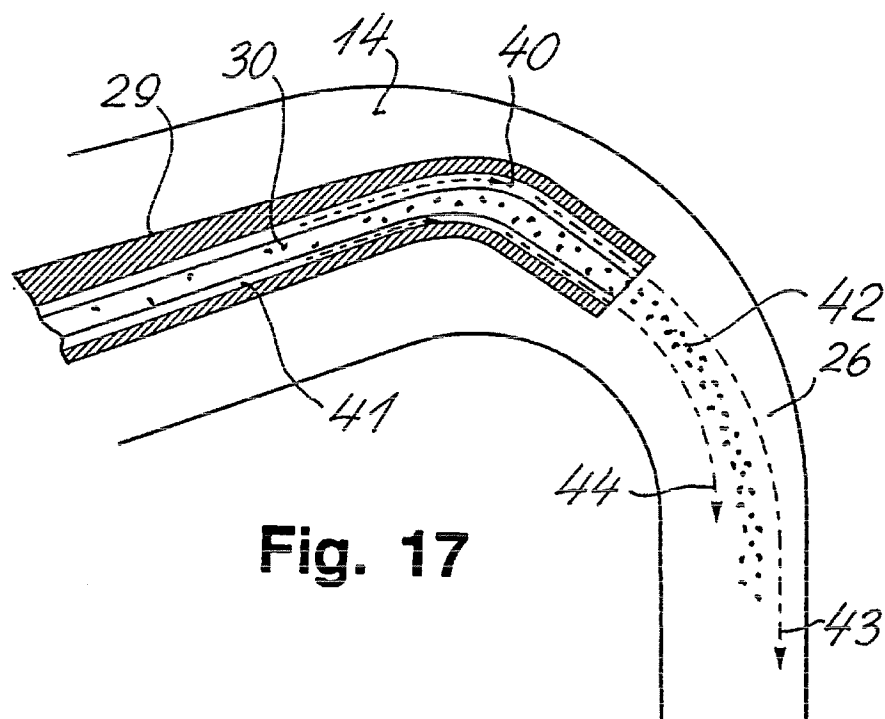
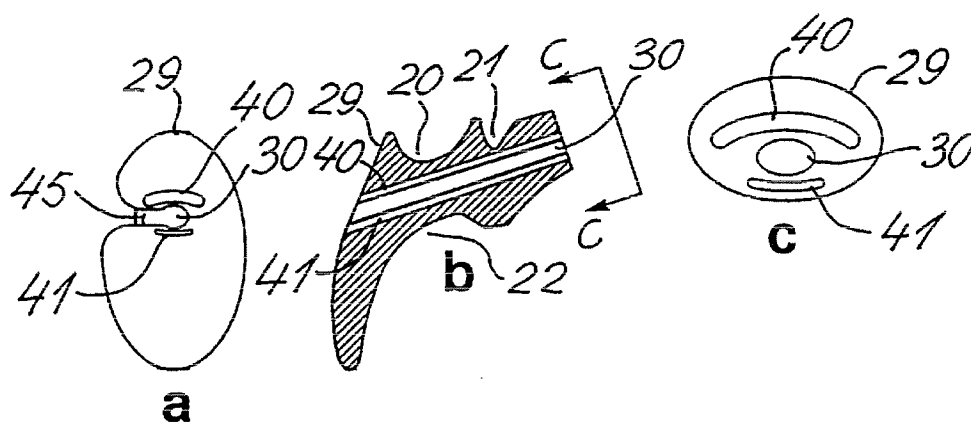


**Fig. 15**

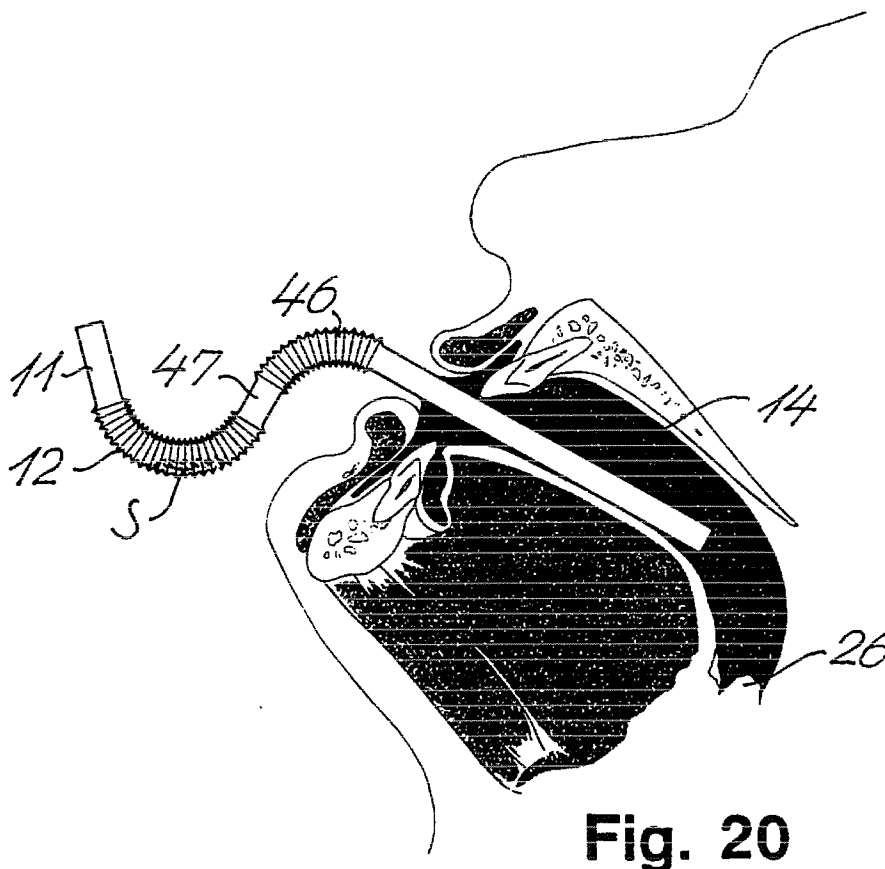
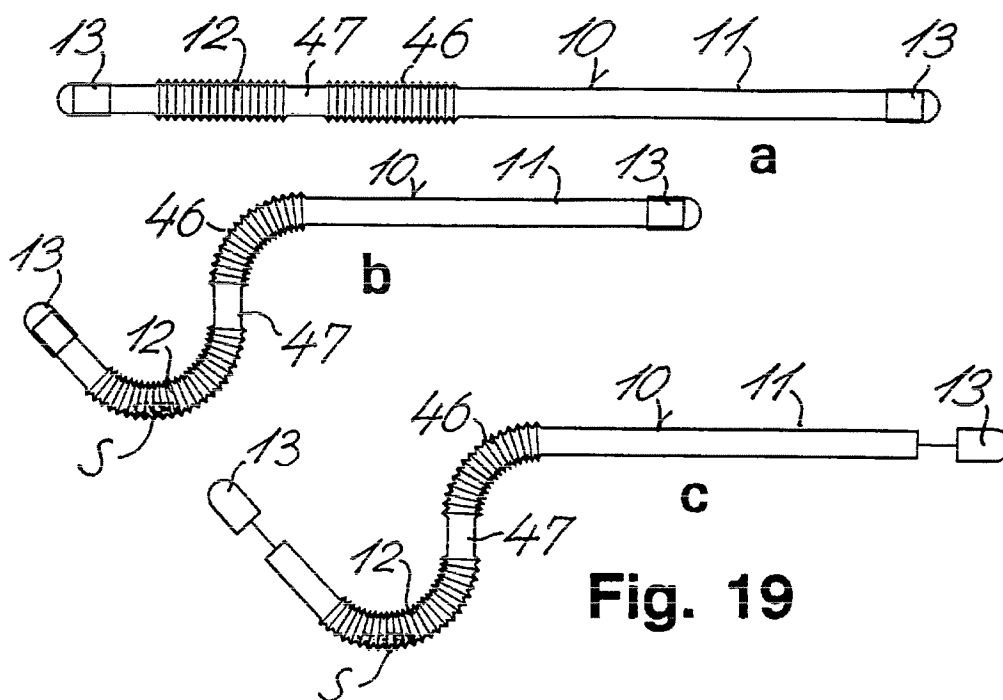


**Fig. 16**

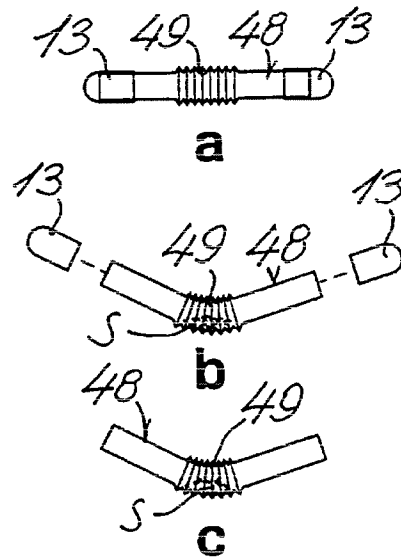
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**Fig. 17****Fig. 18**

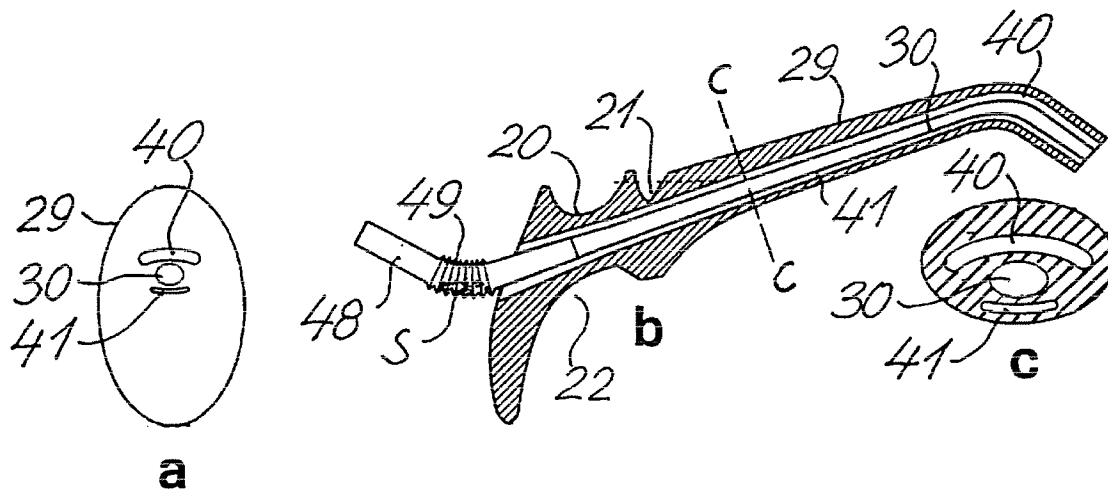
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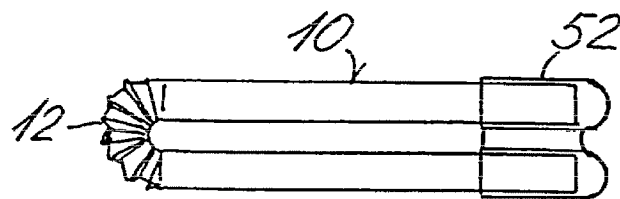
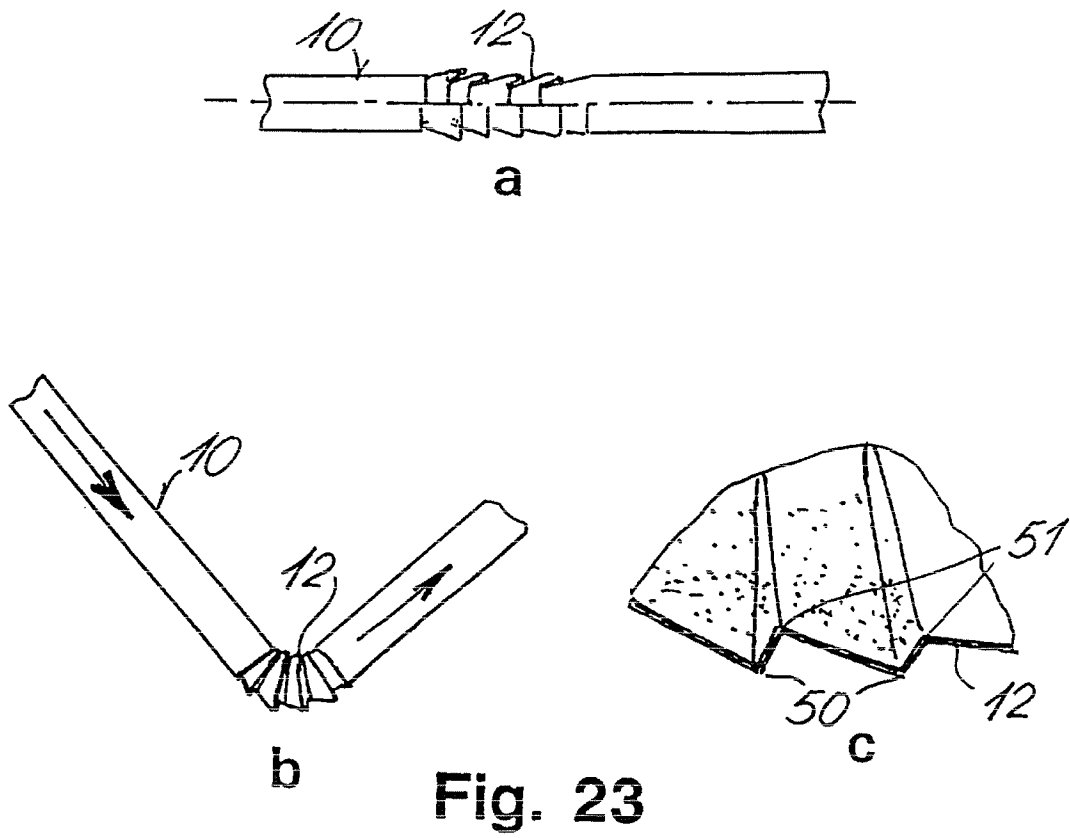
10/11



**Fig. 21**



**Fig. 22**



## INTERNATIONAL SEARCH REPORT

International application No.

PCT/DK 96/00034

## A. CLASSIFICATION OF SUBJECT MATTER

IPC6: A61M 15/00

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC6: A61M

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

SE,DK,FI,NO classes as above

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 2503732 A (C.A. HEISTERKAMP), 11 April 1950 (11.04.50), see whole document  --	1-8,21-30
X	WO 9317728 A1 (AKTIEBOLAGET ASTRA), 16 Sept 1993 (16.09.93), page 12, line 4 - line 19, figure 1, abstract  --	1,3-6,13-14
X	EP 0404454 A1 (FISONS PLC), 27 December 1990 (27.12.90), abstract  --	1,3-6
X	WO 9405358 A1 (MEDIX LIMITED), 17 March 1994 (17.03.94), page 14, line 14 - page 15, line 17, abstract  --	2,4-9,13, 21-26

☒ Further documents are listed in the continuation of Box C.☒ See patent family annex.

- \* Special categories of cited documents:
- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

18 June 1996

Date of mailing of the international search report

19 -06- 1996

Name and mailing address of the ISA/

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# INTERNATIONAL SEARCH REPORT

International application No.

PCT/DK 96/00034

## C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 4265236 A (A.M. PACELLA), 5 May 1981 (05.05.81), column 3, line 23 - line 61, figures 2-4	1,3-13,18-30
Y	--	14-17
X	WO 8901348 A1 (TEIJIN LIMITED), 23 February 1989 (23.02.89), page 5, line 35 - page 6, line 18; page 7, line 7 - line 16, figures 6-7	1,3-13,18-30
Y	--	14-17
Y	US 4148308 A (W.J. SAYER), 10 April 1979 (10.04.79), column 4, line 8 - line 19	14-17
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# INTERNATIONAL SEARCH REPORT

International application No.

PCT/DK 96/00034

## Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:
2. ☐ Claims Nos.:  
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. Claims 1, 3-30 relate to a disposable inhaler comprising a single dose of an active substance.
  11. Claims 2, 4-30 relate to an inhaler with means for supplying a dose into the flow passage and to construction means of the flow passage.
  111. Claims 31-35 relate to a substance supply means for use in connection with an inhaler.
- 
1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
  2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
  3. ☒ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.: 1-30
  4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

☐

The additional search fees were accompanied by the applicant's protest.

☒

No protest accompanied the payment of additional search fees.

# INTERNATIONAL SEARCH REPORT

International application No.

PCT/DK 96/00034

Patent document cited in search report		Publication date	Patent family member(s)		Publication date
US-A-	2503732	11/04/50	NONE		
WO-A1-	9317728	16/09/93	AU-B-	666171	01/02/96
			CA-A-	2131157	16/09/93
			CZ-A-	9402084	18/01/95
			EP-A,A-	0558879	08/09/93
			EP-A-	0629136	21/12/94
			FI-A,D-	944034	02/09/94
			HU-A-	69090	28/08/95
			HU-D-	9402541	00/00/00
			JP-T-	7508184	14/09/95
			NO-A,D-	943211	30/08/94
			NZ-A-	249128	26/01/96
			SK-A-	104994	12/04/95
			ZA-A-	9301520	06/09/93
EP-A1-	0404454	27/12/90	SE-T3-	0404454	
			CA-A-	2019385	21/12/90
			DE-T-	69002800	23/12/93
			ES-T-	2043285	16/12/93
			JP-A-	3037077	18/02/91
			US-A-	5239991	31/08/93
WO-A1-	9405358	17/03/94	NONE		
US-A-	4265236	05/05/81	NONE		
WO-A1-	8901348	23/02/89	AU-B,B-	608895	18/04/91
			AU-A-	2269588	09/03/89
			CA-A-	1329526	17/05/94
			DE-A-	3871131	17/06/92
			EP-A,A,B	0328685	23/08/89
			SE-T3-	0328685	
			JP-T-	2500172	25/01/90
US-A-	4148308	10/04/79	CA-A-	1114702	22/12/81
			CH-A-	624005	15/07/81
			DE-A,A-	2815039	14/12/78
			FR-A,B-	2392649	29/12/78
			GB-A-	1553174	19/09/79
			SE-B,C-	430208	31/10/83
			SE-A-	7803452	01/12/78